#### **IN THE SPECIFICATION:**

## Please replace pages 10-14 ("BRIEF DESCRIPTION OF THE FIGURES") with the attached substitute pages 10-14.

#### Please rewrite the paragraph beginning at page 45, line 2 as follows:

--Sequence of the pGEX and mcg7 (underlined) junction:

pGEX-1

mcg7 (695)

٦١

Sj26...GAA TTC GGC ACG AGC CGA CGG (SEQ ID NO: 20)

Additional amino acids:

Glu Phe Gly Thr Ser (SEQ ID NO: 111).--

#### **IN THE SEQUENCE LISTING:**

Please replace the Sequence Listing of Record with the attached substitute

Sequence Listing.

#### **REMARKS**

Applicants respectfully submit that the specification has been amended to insert the sequence identifiers at pages 10-14 (Brief Description of the Figures) for those sequences disclosed in the figures. The specification has also been amended at page 45 to insert the sequence identifier for the amino acid sequence disclosed at line 2 of page 45. Applicants submit that the foregoing amendment does not introduce new matter.

Applicants have also amended the Sequence Listing of record. More specifically, after SEQ ID NO: 9 on page 74 and before SEQ ID NO: 10 on page 75, there are three sequences that are designated as SEQ ID NO: 7, SEQ ID NO: 8, and SEQ ID NO: 9. Applicants respectfully submit that these sequences were introduced as a result of a clerical error, and have

been deleted from the Sequence Listing as indicated in the attached marked-up copy of the Sequence Listing. In addition, Applicants have reformatted the sequences of SEQ ID NO: 1 to SEQ ID NO: 45 using the PatentIn Version 2.1. Furthermore, Applicants have added sequences designated as SEQ ID NOS: 46-111, which are disclosed in the Figures of the application as originally filed.

Applicants submit that the foregoing amendment to the Sequence Listing does not introduce new matter. Applicants submit herewith a substitute paper and initial computer readable copy of the Sequence Listing, along with a Statement Under 37 C.F.R. §1.821(f), stating that these copies are identical. A copy of the Notice to Comply is also enclosed.

Attached hereto is a marked-up version of the changes made to the specification and claims by the instant amendment. The attached page is captioned "Version with Markings to **Show Changes Made.**"

In view of the foregoing amendments and remarks, it is firmly believed that the subject application is in condition for allowance, which action is earnestly solicited.

Respectfully submitted,

Frank S. DiGiglio

Registration No. 31,346

SCULLY, SCOTT, MURPHY & PRESSER 400 Garden City Plaza Garden City, New York 11530 (516) 742-4343

FSD/XZ:ab

Encls.: Substitute copy of pages 10-14 of the Specification Version with Markings to Show Changes Made Marked-up copy of the Sequence Listing



Serial No:

09/424,458

#### **VERSION WITH MARKINGS TO SHOW CHANGES MADE**

#### **IN THE SPECIFICATION:**

#### Please amend the paragraph beginning at page 45, line 2, as follows:

--Sequence of the pGEX and mcg7 (underlined) junction:

pGEX-1 mcg7 (695)

Sj26...GAA TTC GGC ACG AGC CGA CGG ([SEQ ID NO: 20])

Additional amino acids:

Glu Phe Gly Thr Ser (SEQ ID NO: 111).--

#### Please amend pages 10-14 ("BRIEF DESCRIPTION OF THE FIGURES") as follows:

#### --BRIEF DESCRIPTION OF THE FIGURES

Figure 1 is a representation of the nucleotide sequence ([SEQ ID NO: 2]) and corresponding amino acid sequence ([SEQ ID NO: 3]) of mcg4.

Figure 2 is a representation of the alignment of the human MCG4 amino acid sequence (amino acids: 1-60, SEQ ID NO: 46) (Query) with a translation of a partial murine expressed sequence tag (EST) (SEQ ID NO: 47) (Subject).

Figure 3 is a representation of the alignment of the human MCG4 amino acid sequence (amino acids 7-109; SEQ ID NO: 48) (Query) with a translation of a partial nematode EST (SEQ ID NO: 49) (subject).

Figure 4 is a diagrammatic representation showing a predicted structure of MCG4 where H and C represent histidine and cysteine residues, respectively and X refers to any amino acid residue. Zn [represent] represents zinc atoms.

Figure 5 is a representation of sensitive sequence homology search of related cysteine-

containing motifs in another *Caenorhabditis elegans* protein. Query: amino acids 56-100 (SEQ ID NO: 50). Subject: SEQ ID NO: 51.

Figure 6 is a representation showing that a related cysteine containing motif is present in the GATA-binding transcription factor from *Sacchromyces pombe*. Queries: amino acids 35-58 (SEQ ID NO: 52); amino acids 162-227 (SEQ ID NO: 54). Subjects: amino acids 175-198 (SEQ ID NO: 53); amino acids 441-506 (SEQ ID NO: 55).

Figure 7 is a Northern blot showing expression of *mcg*4 in various cultured human cancer cell lines. Lanes 1-5, respectively, represent the hybridization signal from 15 µg total RNA derived from various human cancer cells lines. Lanes 1-5, respectively, contain RNA from H69 lung carcinoma cells, JAM ovary carcinoma cells, BT20 breast carcinoma cells, HaCat transformed keratinocytes, T24 bladder carcinoma cells.

Figure 8 is a representation of a partial alignment of *mcg*4 with human ESTs AA074703 and AA134788. Queries: nucleotides 446-704 (SEQ ID NO: 56); nucleotides 398-452 (SEQ ID NO: 58); nucleotides 767-810 (SEQ ID NO: 60); nucleotides 731-765 (SEQ ID NO: 62); nucleotides 701-732 (SEQ ID NO: 64); nucleotides 498-687 (SEQ ID NO: 66); nucleotides 398-495 (SEQ ID NO: 68); nucleotides 702-761 (SEQ ID NO: 70). Subjects: nucleotides 49-307 (SEQ ID NO: 57); nucleotides 2-56 (SEQ ID NO: 59); nucleotides 373-416 (SEQ ID NO: 61); nucleotides 336-370 (SEQ ID NO: 63); nucleotides 305-336 (SEQ ID NO: 65); nucleotides 103-292 (SEQ ID NO: 67); nucleotides 2-99 (SEQ ID NO: 69); nucleotides 309-368 (SEQ ID NO: 1).

Figure 9 is a representative of the partial nucleotide sequence alignment between a human (W32939) (SEQ ID NO: 72) and mouse (AA242159) (SEQ ID NO: 73) mcg4-like EST in the putative 5' UTR of the mcg4 cDNA. The putative initiation codon is underlined and the region

upstream represents 5' UTR.

Figure 10 is a representation showing MacVector alignment of MCG4 with forward translations of ESTs AA134788 and AA074703. The nucleotide sequences are shown in Figure 8.

Figure 11 is a diagrammatic representation of the domains of MCG4

zinc finger consensus: CX<sub>2</sub>HX<sub>4</sub>CX<sub>2</sub>CX<sub>4</sub>HX<sub>2</sub>CX<sub>17</sub>CX<sub>2</sub>CX<sub>18</sub>HX<sub>2</sub>CX<sub>18</sub>CX<sub>2</sub>C (SEQ ID NO: 74);

acidic domain consensus: 9/34 amino acids negatively charged, 0/34 positively charged;

basic domain consensus: 13/55 amino acids positively charged, 0/55 negatively charged;

leucine zipper domain consensus: LX<sub>6</sub>LX<sub>6</sub>RX<sub>6</sub>LX<sub>6</sub>L (SEQ ID NO: 75);

alternate "novel" leucine zipper-like motif where leucine would not be aligned along the one surface to an alpha helix domain: (aa261) LX<sub>6</sub>LXLX<sub>6</sub>LXLX<sub>6</sub>L (aa286) (SEQ ID NO: 76).

Figure 12 is a representation showing similarity of MCG7 WITH GEFs of various organisms.

Figure 13(a) is a representation of the nucleotide sequence ([SEQ ID NO: 4]) and corresponding amino acid sequence ([SEQ ID NO: 5]) of mcg7. Nucleotides 183-288 are an alternative spliced exon (shown in lower case).

Figure 13(b) is a representation of the partial nucleotide sequence (§SEQ ID NO: 6]) and corresponding amino acid sequence (§SEQ ID NO: 7]) OF *mcg7* but without the exon shown in Fig. 13(a). Amino acids have been numbered from the first methionine codon (underlined). The cDNA molecules of Fig. 13(a) and Fig. 13(b) differ by the inclusion and exclusion of the exon of nucleotides 183-288.

Figure 14 is a representation showing a comparison between MCG7 (SEQ ID NO: 7) and

a homologue from *Caenorhabditis elegans* (SEQ ID NO: 77) using the BESTFIT algorithm. In the figure, the following sequences are underlined:

#### EF-Hand=PROSITE DATABASE NO. PD0C00018

la nematode DVDEEDEVEDIEF ({SEQ ID NO: 10})

lb human DVDGDGHISQEEF ({SEQ ID NO: 11})

nematode DHDRDGFISQEEF ({SEQ ID NO: 12})

lc human DQNQDGCISREEM ({SEQ ID NO: 13})

nematode DVDMDGQISKDEL (fSEQ ID NO: 14)

#### **GUANINE NT BINDING REGION = BLOCKS DATABASE NO. BL00720B**

2 human HFVHVAEKLLQLQNFNTLMAVVGGLSHSSISRLKETH ({SEQ ID NO: 15})
nematode KFVHVAKHLRKINNFNTLMSVVGGITHSSVARLAKTY ({SEQ ID NO: 16})

#### **DaG-PE BINDING DOMAIN = PROSITE DATABASE NO. PD0C00379**

3 human HNFQESNSLRPVACRHCKALILGIYKQGLKCRACGVNCHKQCKDRLSVEC
({SEQ ID NO: 17})

nematode HNFHETTFLTPTTCNHCNKLLWGILRQGFKCKDCGLAVHSCCKSNAVAEC
({SEQ ID NO: 18})

Figure 15 is a representation of an alignment of human and a partial (5' UTR and partial coding sequence) murine *mcg*7 cDNA (GenBank Acc.No. W71787 and AA237373). The putative initiation codon is underlined. The murine sequence (SEQ ID NO: 78) represents a composite of 2 partial cDNA sequences from the EST database (accession numbers W71787 and AA237373). Nucleotide differences between human and murine sequences are shown in lower case lettering and identical residues are indicated with asterisks.

Figure 16 is a representation of further 5' nucleotide (SEQ ID NO: 79) and corresponding amino acid sequence (SEQ ID NO: 80) human *mcg*7. Nucleotide positions 1-321 were derived from GenBank Acc. No. AC000134 and nucleotides 322 onwards from Figure 13(a). Two inframe initiation codons are underlined. Asterisks denote in-frame stop codons.

Figure 17 is a graphical representation of a GDP release assay. DExperiment #1 (mean of duplicate). ♦ Experiment #2 (mean of duplicates). The exchange reaction contained 36pmols of GST-MCG (N-terminally truncated; encoded by Construct B in Fig. 18) and 1.6-12.8 pmols of recombinant GST-N-Ras.GDP. Reaction time 6 min.

Estimated reaction constants:

 $K_m = 2.1 \mu M$ ,  $V_{max} = 37 \text{ pMol/6min/36pMol [Expt.#1]}$ 

 $K_m = 1.5 \mu M$ ,  $V_{max} = 30.3 p Mol/6 min/36 p Mol[Expt.#2]$ 

Figure 18 depicts various recombinant plasmids containing partial or full-length mcg7.

Figure 19 is a representation of the nucleotide sequence ({SEQ ID NO: 8}) and corresponding amino acid sequence ({SEQ ID NO: 9}) of *mcg18*.

Figure 20 is representation showing that MCG18 (amino acids 35-96, SEQ ID NO: 81] (Query) has partial homology to E. coli DnaJ (amino acids 6-65, SEQ ID NO: 82) (Subject).

Figure 21 is a representation showing that MCG18 has homotology to two *Caenorhabitis* elegans proteins. Queries: amino acids 28-64 (SEQ ID NO: 83); amino acids 71-102 (SEQ ID NO: 85); amino acids 162-203 (SEQ ID NO: 87); amino acids 35-75 (SEQ ID NO: 89); amino acids 75-96 (SEQ ID NO:91); amino acids 141-184 (SEQ ID NO: 93). Subjects: amino acids 22-58 (SEQ ID NO:84); amino acids 64-95 (SEQ ID NO: 86); amino acids 158-199 (SEQ ID NO: 88); amino acids 19-50 (SEQ ID NO: 90); amino acids 71-92 (SEQ ID NO: 92); amino acids

### **IN THE SEQUENCE LISTING:**

Please amend the Sequence Listing of Record as indicated in the attached marked-up copy
of the amended sequence listing.

# Marked-up Copy -57 of the Sequence Listing. SEQUENCE LISTING

Serial No. 09/424,458

DEC 1 8 2001 25

(1) GENERAL INFORMATION:

(APPLICANT: (OTHER THAN US):

The Council of The Queensland Institute of

Medical Research

(US ONLY): HAYWARD Nicholas, SILINS Ginters, GRIMMOND Sean.

GARTSIDE Michael and HANCOCK, John

(ii) TITLE OF INVENTION: A NOVEL GENE AND USES THEREFOR

(iii) NUMBER OF SEQUENCES: 45

(iv) CORRESPONDENCE ADDRESS:

- (A) ADDRESSEE: DAVIES COLLISON CAVE
- (B) STREET: 1 LITTLE COLLINS STREET
- (C) CITY: MELBOURN
- (D) STATE: VICTORIA
- (E) COUNTRY: AUSTRALI
- (F) ZIP: 3000

(v) COMPUTER READABLE FORM:

- (A) MEDIUM TYPE: Floppy disk
- (B) COMPUTER: IBM PC compatible
- (C) OPERATING SYSTEM: PC-DOS/MS-DOS
- (D) SOFTWARE: PatentIn Release #1.0, Version #1.25

(vi) CURRENT APPLICATION DATA:

- (A) APPLICATION NUMBER: PCT INTERNATIONAL
- (B) FILING DATE: 22-MAY-1998
- (C) CLASSIFICATION:

(vii) PRIOR APPLICATION DATA:

- (A) APPLICATION NUMBER: PO6973
- (B) FILING DATE: 23-MAY-1997
- (C) CLASSIFICATION:

(vii) PRIOR/APPLICATION DATA:

- (A) APPLICATION NUMBER: PO6974
- (B) FILING DATE: 23-MAY-1997
- (CYCLASSIFICATION:

(vi) PRIOR APPLICATION DATA:

- (A) APPLICATION NUMBER: PO6972
- (B) FILING DATE: 23 MAY 1997

#### -(C) CLASSIFICATION:

- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: PP1459
  - (B) FILING DATE: 22-JAN-1998
  - (C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: PP/460
  - (B) FILING DATE: 22-JAN-1998
  - (C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:
  - (A) APPLICATION NUMBER: PP1458
  - (B) FILING DATE 22-JAN-1998
  - (C) CLASSIFICATION:
- (viii) ATTORNEY/AGENT INFORMATION:
  - (A) NAME: HUGHES, DR E JOHN L
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  - (B) TELEFAX: +61 3 9254 2770
  - <del>(C) TELEX: AA 31787</del>

(110) Hayward, Nicholes Silinis, Ginters Grimmond, Sean Gartside, Michael Hancock, John

(120) A Novel Gene and Uses Thereof

<130> 13198

<140> 09/424,458

(141 > 1999-11-23

<160> 111

(170 > PaterdIn Ver. 2.1

(2)	INFORMATION	FOR SEO	ID	NO:1:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 8 amino acids
    (B) TYPE: amino acid

  - (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: Peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

Cys Xaa Xaa Cys Xaa Gly Xaa Gly

#### (2) INFORMATION FOR SEQ ID NO:2:

- (i) SEQUENCE CHARACTERISTICS:
  (A) LENGTH: 1242 base pairs

  - (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA
- (ix) FEATURE:
  - (A) NAME/KEY: CDS
  - (B) LOCATION: 30..959

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

TCA	GTAA	ACA	CAGA	GACT	ec c	GATO	GATO	ATC Met	GGG	CTI Leu	TGT Cys	AAG Lys 5	TGC	Pro	AAG Lys	53
AGA Arg	AAG Lys 10	val	ACC	AAC Asn	CTG Leu	TTC Phe 15	Суз	TTC Phe	GAA Glu	CAT	CGG Arg	Val	AAC Asn	GTC Val	TGC Cys	101
GAG Glu 25	CAC His	TGC Cys	CTG Leu	GTA Val	GCC Ala 30	AAT Asn	CAC His	GCC Ala	AAG Lys	TGC Cys 35	ATC Ile	GTC Val	CAG Gln	TCC Ser	TAC Tyr 40	149
CTG Leu	CAA Gln	TGG Trp	CTC Leu	CAA Gln 45	GAT Asp	AGC Ser	GAC Asp	TAC Tyr	AAC Asn 50	CCC Pro	AAT Asn	TGC Cys	CGC Arg	CTG Leu 55	TGC Cys	197
AAC Asn	ATA Ile	CCC Pro	CTG Leu 60	GCC Ala	AGC Ser	CGA Arg	GAG Glu	ACG Thr 65	ACC Thr	CGC Arg	CTT Leu	GTC Val	TGC Cys 70	TAT Tyr	GAT Asp	245
CTC Leu	TTT Phe	CAC His 75	TGG Trp	GCC Ala	TGC Cys	CTC Leu	AAT Asn 80	GAA Glu	CGT Arg	GCT Ala	GCC Ala	CAG Gln 85	CTA Leu	CCC Pro	CGA Arg	293
AAC Asn	ACG Thr 90	GCA Ala	CCT Pro	GCC Ala	GGC Gly	TAT Tyr 95	CAG Gln	TGC Cys	CCC Pro	AGC Ser	TGC Cys 100	AAT Asn	GGC Gly	CCC Pro	ATC Ile	341
TTC Phe 105	CCC Pro	CCA Pro	ACC Thr	AAC Asn	CTG Leu 110	GCT Ala	GGC Gly	CCC Pro	GTG Val	GCC Ala	TCC Ser	GCA Ala	CTG Leu	AGA Arg	GAG Glu	389

115

120

AAG Lys	CTG Leu	GCC Ala	ACA Thr	Val	Asn	TGG Trp	GCC	CGC	GCA Ala 130	Gly	CTG Leu	GGC	CTC Leu	Pro	CTG Leu	437
ATC Ile	GAT Asp	GAG Glu	Val 140	Val	AGC Ser	CCA Pro	GAG Glu	Pro 145	Glu	CCC Pro	CTC Leu	AAC Asn	ACG Thr 150	Ser	GAC Asp	485
TTC Phe	TCT Ser	GAC Asp 155	Trp	TCT Ser	AGT Ser	TTT Phe	AAT Asn 160	GCC Ala	AGC Ser	AGT Ser	ACC Thr	CCT Pro 165	GGA Gly	CCA Pro	GAG Glu	533
GAG Glu	GTA Val 170	GAC Asp	AGC Ser	GCC Ala	TCT Ser	GCT Ala 175	GCC Ala	CCA Pro	GCC Ala	TTC Phe	TAC Tyr 180	AGC Ser	CGA Arg	GCC Ala	CCC	581
CGG Arg 185	CCC Pro	CCA Pro	GCT Ala	TCC Ser	CCA Pro 190	GGC Gly	CGG Arg	CCC	GAG Glu	CAG Gln 195	CAC	ACA Thr	GTG Val	ATC Ile	CAC His 200	629
ATG Met	GCC	AAT Asn	CCT Pro	GAG Glu 205	CCC Pro	TTG Leu	ACT Thr	CAC His	GCC Ala 210	CCT Pro	AGG Arg	AAG Lys	GTG Val	TAT Tyr 215	GAT Asp	677
ACG Thr	CGG Arg	GAT Asp	GAT Asp 220	GAC Asp	CGG Arg	ACA Thr	CCA Pro	GGC Gly 225	CTC Leu	CAT His	GGA Gly	Asp	TGT Cys 230	GAC Asp	GAT Asp	725
GAC Asp	AAG Lys	TAC Tyr 235	CGA Arg	CGT Arg	CGG Arg	CCG Pro	GCC Ala 240	Leu	GGT Gly	TGG Trp	Leu	GCC Ala 245	CGG Arg	CTG Leu	CTA Leu	773
Arg .	AGC Ser 250	CGG Arg	GCT Ala	GIY	Ser	CGG Arg 255	AAG Lys	CGG Arg	CCG Pro	Leu	ACC Thr 260	CTG Leu	CTC Leu	CAG Gln	CGG Arg	821
GCG ( Ala ( 265	GGG Gly	CTG Leu	CTG Leu	Leu	CTC Leu 270	TTG Leu	GGA Gly	CTG Leu	CTG Leu	GGC Gly 275	TTC ( Phe	CTG ( Leu 1	GCC Ala	Leu	CTT Leu 280	869
GCC ( Ala 1	CTC . Leu :	ATG Met	Ser .	CGC Arg 285	CTA ( Leu (	GGC (	CGG ( Arg /	Ala	GCA ( Ala : 290	GCT Ala	GAC A	AGC ( Ser )	lsp	CCC . Pro . 295	AAC Asn	917
CTG ( Leu A	SAC (	Pro :	CTC I Leu I 300	ATG A	AAC ( Asn )	CCT ( Pro 1	lis :	ATC   Ile . 305	CGC ( Arg \	GTG ( Val (	GGC (	Pro S	rcc : Ser 310	TGA		962
cccc	CTT	GC T	rgrg	CTA	GCC	CAGC	CTAG	GAT	GTGGG	SŤT (	CTGTG	GAGG	A G	AGGC	GGGT	1022
ATGG	GGA	GG C	TGAG	GCA	CTC	CTTC	CTG	CCC	CTCTC	ccc :	CAAC	CCTA	A G	ACAC	raaga	1082
															CCTG	1142
GTCA	AGC	AT TO	rgtci	TGAC	TTC	CTTI	CTC	CCG	GTC1	rcc z	AGCCI	CCGA	c cc	CTC	cccc	1202
TGAA	GGA	C TO	GCAC	GTGC	AA.	<b>LAAT</b>	CAA	CAAC	TTTA	TT					,	1242

### (2) INFORMATION FOR SEQ ID NO:3:

- (i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 310 amino acids(B) TYPE: amino acid(D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Met Gly Leu Cys Lys Cys Pro Lys Arg Lys Val Thr Asn Leu Phe Cys
1 10 15

Phe Glu His Arg Val Asn Val Cys Glu His Cys Leu Val Ala Asn His 20 25 30

Ala Lys Cys Ile Val Gln Ser Tyr Leu Gln Trp Leu Gln Asp Ser Asp
35 40 45

Tyr Asn Pro Asn Cys Arg Leu Cys Asn Ile Pro Leu Ala Ser Arg Glu
50 60

Thr Thr Arg Leu Val Cys Tyr Asp Leu Phe His Trp Ala Cys Leu Asn 65 70 75 80

Glu Arg Ala Ala Gln Leu Pro Arg Asn Thr Ala Pro Ala Gly Tyr Gln 85 90 95

Cys Pro Ser Cys Asn Gly Pro Ile Phe Pro Pro Thr Asn Leu Ala Gly 100 105 110

Pro Val Ala Ser Ala Leu Arg Glu Lys Leu Ala Thr Val Asn Trp Ala 115 120 125

Arg Ala Gly Leu Gly Leu Pro Leu Ile Asp Glu Val Val Ser Pro Glu 130 135 140

Pro Glu Pro Leu Asn Thr Ser Asp Phe Ser Asp Trp Ser Ser Phe Asn 145 150 155 160

Ala Ser Ser Thr Pro Gly Pro Glu Glu Val Asp Ser Ala Ser Ala Ala 165 170 175

Pro Ala Phe Tyr Ser Arg Ala Pro Arg Pro Pro Ala Ser Pro Gly Arg 180 185 190

Pro Glu Gln His Thr Val Ile His Met Gly Asn Pro Glu Pro Leu Thr 195 200 205

His Ala Pro Arg Lys Val Tyr Asp Thr Arg Asp Asp Asp Arg Thr Pro 210 215 220

Gly Leu His Gly Asp Cys Asp Asp Lys Tyr Arg Arg Arg Pro Ala 225 230 235 240

Leu Gly Trp Leu Ala Arg Leu Leu Arg Ser Arg Ala Gly Ser Arg Lys
245 250 255

Arg Pro Leu Thr Leu Leu Gln Arg Ala Gly Leu Leu Leu Leu Gly 260 265 270

Leu Leu Gly Phe Leu Ala Leu Leu Ala Leu Met Ser Arg Leu Gly Arg 275 280 285

Ala Ala Ala Asp Ser Asp Pro Asn Leu Asp Pro Leu Met Asn Pro His 290 295 300

Ile Arg Val Gly Pro Ser 305 310

#### (2) INFORMATION FOR SEQ ID NO:4:

(i) SEQUENCE CHARACTERISTICS:

(A)	LENGTH:	2415	base	pairs
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(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

#### (ii) MOLECULE TYPE: DNA

#### (ix) FEATURE:

(A) NAME/KEY: CDS
(B) LOCATION: 3...2188 2186

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

								_								
CG	ATT Ile 1	TCA Ser	TTC Phe	CTC Leu	GCT Ala 5	CCC Pro	CAC His	AGG Arg	TCC	CTC Leu 10	TCC Ser	CCA Pro	AAA Lys	TAT Tyr	TCC Ser 15	47
CAT	CTT Lev	GTC Val	CTA Leu	GCC Ala 20	His	CCC Pro	CCA Pro	GAC Asp	TAT Tyr 25	Lev	Lys	GA( Asp	C.CAC	G CTO Leu 30	TCC Ser	95
CCA Pro	CGC Arg	CCC Pro	CGA Arg 35	Pro	CCA Pro	CTA Leu	GGC	CTG Leu 40	Cys	CAC His	CCG Pro	Leu	CCT Pro	Ala	GGA Gly	143
AGA Arg	CGC	CCG Pro -50	Val	CCG	GGC	CGG Arg	GTT Val 55	AGC Ser	CCC Pro	ATG Met	GGA Gly	ACG Thr 60	Gln	CGC Arg	CTG Leu	191
TGT Cys	GGC Gly 65	Arg	GGG Gly	ACT Thr	CAA Gln	GGC Gly 70	TGG Trp	CCT Pro	GGC Gly	TCA Ser	AGT Ser 75	GAA Glu	CAG Gln	CAC	GTC Val	239
CAG Gln 80	GAG Glu	GCG Ala	ACC	TCG Ser	TCC Ser 85	GCG Ala	GGT Gly	TTG Leu	CAT His	TCT Ser 90	GGG	GTG Val	GAC Asp	GAG Glu	CTG Leu 95	287
GGG Gly	GTT Val	CGG	TCC Ser	GAG Glu 100	CCC Pro	GGT Gly	GGG Gly	AGG Arg	CTC Leu 105	CCG Pro	GAG Glu	CGC	AGC Ser	CTG Leu 110	GGC	335
CCA Pro	GCC Ala	CAC His	CCC Pro 115	GCG Ala	CCG Pro	GCG Ala	GCC Ala	ATG Met 120	GCA Ala	GGC Gly	ACC Thr	CTG Leu	GAC Asp 125	CTG Leu	GAC Asp	383
AAG Lys	GGC Gly	TGC Cys 130	ACG Thr	GTG Val	GAG Glu	GAG Glu	CTG Leu 135	CTC Leu	CGC Arg	GGG Gly	TGC Cys	ATC Ile 140	GAA Glu	GCC Ala	TTC Phe	431
GAT Asp	GAC Asp 145	TCC Ser	Gly	AAG Lys	GTG Val	CGG Arg 150	GAC Asp	CCG Pro	CAG Gln	CTG Leu	GTG Val 155	CGC Arg	ATG Met	TTC Phe	CTC Leu	479
ATG Met 160	ATG Met	CAC His	CCC Pro	TGG Trp	TAC Tyr 165	ATC Ile	CCC Pro	TCC Ser	Ser	CAG Gln 170	CTG Leu	GCG Ala	GCC Ala	AAG Lys	CTG Leu 175	527
CTC Leu	CAC His	ATC Ile	Tyr	CAA Gln 180	CAA Gln	TCC Ser	CGG Arg	Lys	GAC Asp 185	AAC Asn	TCC Ser	AAT Asn	TCC Ser	CTG Leu 190	CAG Gln	575
GTG Val	AAA Lys	Thr	TGC Cys 195	CAC His	CTG Leu	GTC .	Arg	TAC Tyr 200	TGG Trp	ATC Ile	TCC Ser	Ala	TTC Phe 205	CCA Pro	GCG Ala	623

GAC Glu	TTI Phe	GAG Asj 210	p Lei	G AAC 1 Asr	CCC	G GAG	Leu 215	Ala	r GAG	G CA	G AT	C AA e Ly 22	s Gl	G CT	G AAG u Lys	67	L
		Le					Asn					r Se			C GAC e Asp	719	}
	qeA :					Tyr					y Gli				G CGG Arg 255	767	,
			. Gly		Lys					Ser					CAC His	815	,
				Glu					Leu					тут	CGC Arg	863	
			Lys			Phe							• Val		CAT His	911	
												Ile			TTC Phe	959	
AAC Asn 320	AGC Ser	GTC Val	TCA Ser	CAG Gln	TGG Trp 325	GTG Val	CAG Gln	CTC Leu	ATG Met	ATC Ile 330	CTC Leu	AGC Ser	AAA Lys	CCC	ACA Thr 335	1007	
GCC Ala	CCG Pro	CAG Gln	CGG Arg	GCC Ala 340	CTG Leu	GTC Val	ATC Ile	ACA Thr	CAC His 345	TTT	GTC Val	CAC	GTG Val	GCG Ala 350	GAG Glu	1055	
AAG Lys	CTG Leu	CTA Leu	CAG Gln 355	CTG Leu	CAG Gln	AAC Asn	Phe	AAC Asn 360	ACG Thr	CTG Leu	ATG Met	GCA Ala	GTG Val 365	GTC Val	Gly	1103	
GĠC Gly	CTG Leu	AGC Ser 370	CAC His	AGC Ser	TCC Ser	ATC Ile	TCC Ser 375	CGC Arg	CTC Leu	AAG Lys	GAG Glu	ACC Thr 380	CAC His	AGC Ser	CAC His	1151	
GTT Val	AGC Ser 385	CCT Pro	GAG Glu	ACC Thr	Ile	AAG Lys 390	CTC Leu	TGG Trp	GAG Glu	GGT Gly	CTC Leu 395	ACG Thr	GAA Glu	CTA Leu	GTG Val	1199	
ACG Thr 400	GCG Ala	ACA Thr	GGC	Asn	TAT Tyr 405	GGC A	AAC ' Asn '	TAC Tyr	CGG Arg	CGT Arg 410	CGG Arg	CTG Leu	GCA Ala	GCC Ala	TGT Cys 415	1247	
GTG Val	GGC Gly	TTC Phe	Arg	TTC Phe 420	CCG . Pro	ATC (	CTG ( Leu (	Gly	GTG Val 425	His	CTC Leu	AAG Lys	GAC Asp	CTG Leu 430	GTG Val	1295	
GCC Ala	CTG Leu	CAG Gln	CTG Leu 435	GCA   Ala	CTG Leu	CCT ( Pro )	Asp :	rgg Frp 440	CTG Leu	GAC Asp	CCA Pro	Ala	CGG Arg 445	ACC Thr	CGG Arg	1343	
CTC Leu	Asn (	GGG Gly 450	GCC . Ala	AAG . Lys i	ATG I	AAG ( Lys (	CAG ( Gln I	CTC ( Leu	TTT . Phe	AGC Ser	ATC Ile	CTG Leu 460	GAG Glu	GAG Glu	CTG Leu	1391	
Ala :	ATG ( Met 1	GTG Val	ACC .	AGC (	Leu /	CGG C Arg I 170	CA C	CA (	GTA (	Gln	GCC Ala 475	AAC Asn	CCC Pro	GAC Asp	CTG Leu	1439	

CTC Lev 480	ı Se	C CT	G CT	C ACC	G GT0 r Val 485	l Ser	CTC	GA:	r CAG	G TA:	r Gli	G ACC	G GAG	GAT Asp	GAG Glu 495	1487	
CT( Lev	TAI Ty:	C CA	G CTO	Ser 500	Lev	G CAG	Arg	GAC Glu	Pro 505	Arg	Ser	C AAG Lys	TCC Ser	Ser 510	Pro	1535	
ACC	: AG	c CCC	C ACC o Thi 515	: Ser	TGC Cys	ACC Thr	CCA Pro	CCA Pro 520	Pro	CGG Arg	CCC Pro	CCG Pro	GTA Val 525	CTG Leu	GAG Glu	1583	
GAG Glu	TGC	ACC Thi	r Ser	GCT Ala	GCC Ala	AAA Lys	CCC Pro 535	Lys	CTG Lev	GAT Asp	CAG Gln	GCC Ala 540	CTC Leu	GTG Val	GTG Val	1631	
GAG Glu	CAC His 545	: Ile	GAG Glu	AAG Lys	ATG Met	GTG Val 550	GAG Glu	TCT Ser	GTG Val	TTC Phe	CGG Arg 555	Asn	TTT Phe	GAC Asp	GTC Val	1679	
GAT Asp 560	Gly	GA7 Asp	GGC Gly	CAC	ATC Ile 565	TCA Ser	CAG Gln	GAA Glu	GAA Glu	TTC Phe 570	CAG Gln	ATC Ile	ATC Ile	CGT Arg	GGG Gly 575	1727	
AAC Asn	TTC Phe	Pro	TAC Tyr	CTC Leu 580	Ser	GCC Ala	TTT	GGG Gly	GAC Asp 585	CTC Leu	GAC Asp	CAG Gln	AAC Asn	CAG Gln 590	GAT Asp	1775	
GGC Gly	TGC Cys	ATC Ile	AGC Ser 595	AGG Arg	GAG Glu	GAG Glu	ATG Met	GTT Val 600	TCC Ser	TAT Tyr	TTC	CTG Leu	CGC Arg 605	TCC Ser	AGC Ser	1823	
TCT	GTG Val	TTG Leu 610	Gly	GJA GGG	CGC Arg	ATG Met	GGC Gly 615	TTC Phe	GTA Val	CAC His	AAC Asn	TTC Phe 620	CAG Gln	GAG Glu	AGC Ser	1871	
AAC Asn	TCC Ser 625	TTG Leu	CGC Arg	CCC Pro	GTC Val	GCC Ala 630	TGC Cys	CGC Arg	CAC His	TGC Cys	AAA Lys 635	GCC Ala	CTG Leu	ATC Ile	CTG Leu	1919	
GGC Gly 640	ATC Ile	TAC Tyr	AAG Lys	CAG Gln	GGC Gly 645	CTC	AAA Lys	TGC Cys	CGA Arg	GCC Ala 650	TGT Cys	GGA Gly	GTG . Val .	Asn (	TGC Cys 655	1967	
CAC His	AAG Lys	CAG Gln	TGC Cys	AAG Lys 660	GAT Asp	CGC (	CTG ( Leu	Ser	GTT Val 665	GAG Glu	TGT Cys	CGG Arg	CGC A	AGG ( Arg ) 570	GCC Ala	2015	
CAG Gln	AGT Ser	GTG Val	AGC Ser 675	Leu	GAG ( Glu (	GGG '	Ser 1	GCA Ala 680	CCC Pro	TCA Ser	CCC Pro	Ser :	CCC A Pro N 685	ATG ( Met H	CAC His	2063	
AGC Ser	urs	CAT His 690	CAC His	CGC (	GCC ' Ala 1	Phe S	AGC : Ser 1 595	TTC Phe	TCT Ser	CTG ( Leu )	Pro A	CGC ( Arg 1 700	CCT C	GC A	.rg	2111	
ura ,	GGC Gly 705	TCC Ser	AGG Arg	CCT (	Pro (	GAG A Glu 1 710	ATC (	CGT ( Arg (	GAG ( Glu (	Glu	GAG ( Glu \ 715	GTA ( Val (	CAG A	CG G	TG al	2159	
GAG ( Glu 1 720	GAT (	GJA GGG	GTG (	Phe A	GAC / Asp 1 725	ATC C	r oa: I eil	rrg :	TA A	raga:	rgc T	G TGC	GTTGG	ATC		2208	
AAGG!	ACTC.	AT T	CCTG	CTTC	GAG	AAAA	TAC	TTC	AACC	AGA (	CAGO	GAGC	C TG	GGGG	TGTC	2268	
GGGG	AGG.	AG G	CTGG	GATO	GGG	GTGG	GAT	atga	AGGGT	rgg c	ATGO	AGCT	G AG	GGCA	GGGC	2328	

2388

2415

CAGGGCTGGT GTCCCTAAGG TTGTACAGAC TCTTGTGAAT ATTTGTATTT TCCAGATGGA ATAAAAAGGC CCGTGTAATT AACCTTC (2) INFORMATION FOR SEQ ID NO:5: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 728 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (xi) SEQUENCE DESCRIPTION: SEQ ID NO:5: Ile Ser Phe Leu Ala Pro His Arg Ser Leu Ser Pro Lys Tyr Ser His Leu Val Leu Ala His Pro Pro Asp Tyr Leu Lys Asp Gln Leu Ser Pro Arg Pro Arg Pro Pro Leu Gly Leu Cys His Pro Leu Pro Ala Gly Arg Arg Pro Val Pro Gly Arg Val Ser Pro Met Gly Thr Gln Arg Leu Cys Gly Arg Gly Thr Gln Gly Trp Pro Gly Ser Ser Glu Gln His Val Gln Glu Ala Thr Ser Ser Ala Gly Leu His Ser Gly Val Asp Glu Leu Gly Val Arg Ser Glu Pro Gly Gly Arg Leu Pro Glu Arg Ser Leu Gly Pro Ala His Pro Ala Pro Ala Ala Met Ala Gly Thr Leu Asp Leu Asp Lys Gly Cys Thr Val Glu Glu Leu Leu Arg Gly Cys Ile Glu Ala Phe Asp Asp Ser Gly Lys Val Arg Asp Pro Gln Leu Val Arg Met Phe Leu Met Met His Pro Trp Tyr Ile Pro Ser Ser Gln Leu Ala Ala Lys Leu Leu His Ile Tyr Gln Gln Ser Arg Lys Asp Asn Ser Asn Ser Leu Gln Val Lys Thr Cys His Leu Val Arg Tyr Trp Ile Ser Ala Phe Pro Ala Glu Phe Asp Leu Asn Pro Glu Leu Ala Glu Gln Ile Lys Glu Leu Lys Ala Leu Leu Asp Gin Glu Gly Asn Arg Arg His Ser Ser Len Ile Asp Ile

Asp Ser Val Pro Thr Tyr Lys Trp Lys Arg Gln Val Thr Gln Arg Asn

Pro Val Gly Gln Lys Lys Arg Lys Met Ser Leu Leu Phe Asp His Leu

Glu Pro Met Glu Leu Ala Glu His Leu Thr Tyr Leu Glu Tyr Arg Ser 275 280 285

Phe Cys Lys Ile Leu Phe Gln Asp Tyr His Ser Phe Val Thr His Gly 290 295 300

Cys Thr Val Asp Asn Pro Val Leu Glu Arg Phe Ile Ser Leu Phe Asn 305 310 315 320

Ser Val Ser Gln Trp Val Gln Leu Met Ile Leu Ser Lys Pro Thr Ala 325 330 335

Pro Gln Arg Ala Leu Val Ile Thr His Phe Val His Val Ala Glu Lys
340 345 350

Leu Leu Gln Leu Gln Asn Phe Asn Thr Leu Met Ala Val Val Gly Gly 355 360 365

Leu Ser His Ser Ser Ile Ser Arg Leu Lys Glu Thr His Ser His Val 370 375 380

Ser Pro Glu Thr Ile Lys Leu Trp Glu Gly Leu Thr Glu Leu Val Thr 385 390 395 400

Ala Thr Gly Asn Tyr Gly Asn Tyr Arg Arg Arg Leu Ala Ala Cys Val405 415

Gly Phe Arg Phe Pro Ile Leu Gly Val His Leu Lys Asp Leu Val Ala 420 425 430

Leu Gln Leu Ala Leu Pro Asp Trp Leu Asp Pro Ala Arg Thr Arg Leu
435 440 445

Asn Gly Ala Lys Met Lys Gln Leu Phe Ser Ile Leu Glu Glu Leu Ala 450 455 460

Met Val Thr Ser Leu Arg Pro Pro Val Gln Ala Asn Pro Asp Leu Leu465470470475

Ser Leu Leu Thr Val Ser Leu Asp Gln Tyr Gln Thr Glu Asp Glu Leu 485 490 495

Tyr Gln Leu Ser Leu Gln Arg Glu Pro Arg Ser Lys Ser Ser Pro Thr 500 505 510

Ser Pro Thr Ser Cys Thr Pro Pro Pro Arg Pro Pro Val Leu Glu Glu 515 520 525

Trp Thr Ser Ala Ala Lys Pro Lys Leu Asp Gln Ala Leu Val Val Glu 530 535 540

His Ile Glu Lys Met Val Glu Ser Val Phe Arg Asn Phe Asp Val Asp 545 550 555 560

Gly Asp Gly His Ile Ser Gln Glu Glu Phe Gln Ile Ile Arg Gly Asn 565 570 575

Phe Pro Tyr Leu Ser Ala Phe Gly Asp Leu Asp Gln Asn Gln Asp Gly 580 585 590

Cys Ile Ser Arg Glu Glu Met Val Ser Tyr Phe Leu Arg Ser Ser Ser 595 600 605

Val Leu Gly Gly Arg Met Gly Phe Val His Asn Phe Gln Glu Ser Asn 610 620

Ser Leu Arg Pro Val Ala Cys Arg His Cys Lys Ala Leu Ile Leu Gly

625					630	Ò				63	5					640		
Ile	Tyr	Lys	Gln	Gly 645		Lys	Суз	Arg	g Al 65		s G	ly (	/al	Asn	Cys 655			
Lys	Gln	Cys	660		Arg	Leu	Ser	Va: 665		u Cy	s A	rg A	lrg	Arg 670	Ala	Gln		
Ser	Val	Ser 675		Glu	Gly	Ser	Ala 680		Se.	r Pr	o S		Pro 185	Met	His	Ser		-
His	His 690	His	Arg	Ala	Phe	Ser 695	Phe	Sex	Le	u Pr	o A:		ro	Gly	Arg	Arģ	,	
Gly 705	Ser	Arg	Pro	Pro	Glu 710	Ile	Arg	Glu	ı Glı	Gl: 71:		al G	ln	Thr	Val	Glu 720		
Asp	Gly	Val	Phe	Asp 725	Ile	His	Leu	٠.										
(2)	INFO	RMA	тіои	FOR	SEQ	ID 1	10 : 6	:								٠.		
-	(i)	() ()	A) LE B) TY C) ST	ENGT: (PE: (RANI	i: 2: nuc: EDNI	CTERI 309 h leic ESS: line	ase acio sino	pai i	rs									
	(ii)	MOI	LECUI	E TY	PE:	DNA												
	(ix)	(2	ATURE A) NA B) LC	ME/X		CDS 254.	. 2018	33° 2	080						•			
	(xi)	SEC	UENC	E DE	SCRI	PTIO	N: 5	EQ 1	ID N	0:6:		•					. `	
CGAT	rtca'	TT C	CTCG	СТСС	C ·CA	CAGG	TCCC	TCI	rccc	CAAA	AT	ATTC	:CC#	AT C	TTGT	CCTAG		60
CCCAT	rccc	CC A	GACT	ATCT	C AA	GGAC	CAGO	TGI	rccc	CACG	CCC	ccc	ACC	T C	CACT.	AGGCC	1	120
TGTG	CAC	CC G	CTGC	CTGC.	A GG	AAGA	cGCC	CGG	TCC	CGGG	ccc	GGT	TAC	sc co	CĂT	GGGAA	1	180
CGGGG	TTC	GG T	CCGA	GCCC	G GT	GGGA	GGCT	ccc	GGA	SCGC	AGO	CTG	GGC	c cz	AGCC	CACCC	2	240
CGCGC	CGG	CG G	CC A	TG G et A	CA G la G	GC AG	CC C	TG G eu A 5	SAC (	CTG ( Leu )	GAC Asp	AAG Lys	Gl	C T( Y C) 0	GC AC	IG nr	2	289
GTG G Val G	SAG (	SAG Slu 15	CTG ( Leu )	CTC ( Leu )	CGC ( Arg (	GGG ? Gly (	rgc Ys 20	ATC Ile	GAA Glu	GCC Ala	Phe	GA As	ρA	AC 1	CC ( Ser (	GG Gly	. 3	37
AAG G Lys V	TG ( al / 30	CGG Arg	GAC (	CCG ( Pro (	CAG ( Gln )	CTG ( Leu \ 35	GTG ( /al .	CGC Arg	ATG Met	TTC Phe	CTC Leu 40	Me	G A t M	TG C	CAC C	,ro ;cc	3	85
TGG T Trp T 45	AC A	ATC (	CCC ? Pro !	rcc :	CT ( Ser ( 50	CAG ( Gln I	eu i	GCG Ala	GCC Ala	AAG Lys 55	CTG Leu	CT(	C C	AC A is I	TC T le T	AC Yr 60	4	33
CAA C Gln G	AA 1	cc ( er	CGG <i>I</i> Arg I	AAG ( Lys A 65	SAC A	AAC T Asn S	cc i Ser i	AAT Asn	TCC Ser 70	CTG Leu	CAG Gln	GT( Va)	G A.	ys T	CG Thr C	GC ys	4	81
CAC C	TG G	TC I	AGG 1	TAC 1	GG 1	ATC I	cc c	SCC 1	TTC	CCA	GCG	GAC	3 T'	TT G	AC T	TG	5	29

нi	s Le	u Va		g Ty	r Tr	p Il	e Ser	Ala 85		e Pr	o Al	La G		he A 90	.sp	Leu		
AA As:	c cc n Pr	o Gl	G TT u Le 5	G GC u Al	T GAG	G CA( u Gli	TATO TILE 100	Lys	G GA	G CT	G AA u Ly	AG GC 's Al	la Le	rg c	TA eu	GAC Asp		577
Gl	n Gl	u Gl O	y As	n Ar	g Ar	g His 115	-	Ser	Lei	ı 11	e As 12	p I1 0	e As	p Se	er	Val		625
Pro 125	Th:	r Ty.	r Ly.	s Tr	130	s Arg	G CAG	Val	Thr	135	n Ar	g As	n Pr	o Va	al	Gly 140		673
Gli	AA Lys	A AAG E Ly:	G CGG s Arg	C AAG g Ly: 14:	Met	S TCC	CTG Leu	TTG Leu	Phe 150	: As	CAC Hi	C CT s Le	G GA	.G CC u Pr 15	0	ATG Met	1	721
GA0	CTC Lev	G GCC	G GAG G Glu 160	ı His	CTC Leu	ACC Thr	TAC	TTG Leu 165	GAG Glu	TAI Tyr	CGC Arg	C TC g Se	C TT r Ph 17	e Cy	C .	AAG Lys		769
Ile	Lev	175	e Glr	a Asp	Tyr	His	AGT Ser 180	Phe	Val	Thr	His	5 Gl; 18	у Су 5	s Th	r'	Val		817
Asp	190	Pro	Val	. Leu	Glu	Arg 195	TTC Phe	Ile	Ser	Leu	200	e Ası )	n Se:	r Va	1 :	Ser		865
G1n 205	Trp	Val	Gln	Leu	Met 210	Ile	CTC Leu	Ser	Lys	Pro 215	Thr	Ala	a Pro	Gl:	n j	rg 20		913
Ala	Leu	Val	Ile	Thr 225	His	Phe	GTC Val	His	Val 230	Ala	Glu	Lys	Lei	Let 235	u G 5	ln		961
ren	GIn	Asn	Phe 240	Asn	Thr	Leu		Ala 245	Val	Val	Gly	Gly	Leu 250	Ser	H	is		1009
ser	ser	255	Ser	Arg	Leu	Lys	GAG / Glu ' 260	Thr	His	Ser	His	Val 265	Ser	Pro	G	lu		L057
ınr.	270	Lys	Leu	Trp	Glu	Gly 275		Thr (	Glu	Leu	Val 280	Thr	Ala	Thr	G	ly	1	105
285	ıyr	GIĀ	Asn	Tyr	Arg 290	Arg	CGG ( Arg I	Seu 1	Ala	Ala 295	Cys	Val	Gly	Phe	3 (	rg 00	1	.153
rne	Pro	TIE	Leu	305 Gly	Val	His :	CTC A Leu I	ys /	Asp 310	Leu	Val	Ala	Leu	Gln 315	Le	eu	1	201
Ala	Leu.	Pro	320	Trp	Leu .	Asp :		11a A 125	Arg '	Thr	Arg	Leu	Asn 330	Gly	A.	la .	1	249
AAG Lys	met	AAG Lys 335	CAG Gln	CTC Leu	TTT :	Ser :	ATC C Ile L 340	rG C	GAG ( Glu (	GAG Glu	Leu	GCC Ala 345	ATG Met	GTG Val	AC Tì	ic ir	1	297

AGC Ser	CTG Leu 350	Arg	CCA Pro	-CCA	GTA Val	CAG Gln 355	Ala	AAC	CCC Pro	GAC Asp	CTG Leu 360	Leu	AGC Ser	CTC	CTC Leu	1345
ACG Thr 365	Val	TCT Ser	CTG Leu	GAT Asp	CAG Gln 370	Tyr	CAG Gln	ACC	GAG Glu	GAT Asp 375	Glu	CTG Lev	TAC	CAC Glr	CTG Leu 380	. 1393
TCC	CTG Leu	CAG Gln	CGG Arg	GAG Glu 385	Pro	CGC	TCC Ser	AAG Lys	Ser 390	Ser	CCA Pro	ACC	AGC Ser	Pro 395	'ACG Thr	1441
									Val					Thr	TCG Ser	1489
						GAT Asp										1537
						TTC Phe 435										1585
						TTC Phe	_	_	_		_					1633
						CTC Leu										1681
						TAT Tyr			Arg							1729
						CAC His										1777
						TGC Cys 515										1825
						GCC Ala										1873
	_	_	_	_		GAG Glu	_	_	_				_			1921
						TCA Ser										1969
						CTG Leu										2017
						GAG Glu 595				Thr						2065
			CAC His		TAAT 610	'AGAT	GC T	GTGG	TTGG	A TC	AAGG.	ACTC	ATT	CCTG	CCT	2120

TTAACCTTC		•				2309
GGTTGTACAG	ACTCTTGTGA	ATATTTGTAT	TTTCCAGATG	GAATAAAAAG	GCCCGTGTAA	2300
TGGGGGTGGG	ATATGAGGGT	GGCATGCAGC	TGAGGGCAGG	GCCAGGGCTG	GTGTCCCTAA	2240
TGGAGAAAAT	ACTTCAACCA	GAGCAGGGAG	CCTGGGGGTG	TCGGGGCAGG	AGGCTGGGGA	2180

#### (2) INFORMATION FOR SEQ ID NO:7:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 609 amino acids
  - (B) TYPE: amino acid
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:7: Met Ala Gly Thr Leu Asp Leu Asp Lys Gly Cys Thr Val Glu Glu Leu Leu Arg Gly Cys Ile Glu Ala Phe Asp Asp Ser Gly Lys Val Arg Asp Pro Gln Leu Val Arg Met Phe Leu Met Met His Pro Trp Tyr Ile Pro Ser Ser Gln Leu Ala Ala Lys Leu Leu His Ile Tyr Gln Gln Ser Arg 50 55 60 Lys Asp Asn Ser Asn Ser Leu Gln Val Lys Thr Cys His Leu Val Arg Tyr Trp Ile Ser Ala Phe Pro Ala Glu Phe Asp Leu Asn Pro Glu Leu Ala Glu Gln Ile Lys Glu Leu Lys Ala Leu Leu Asp Gln Glu Gly Asn Arg Arg His Ser Ser Leu Ile Asp Ile Asp Ser Val Pro Thr Tyr Lys

Trp Lys Arg Gln Val Thr Gln Arg Asn Pro Val Gly Gln Lys Lys Arg

Lys Met Ser Leu Leu Phe Asp His Leu Glu Pro Met Glu Leu Ala Glu 150 155

Kis Leu Thr Tyr Leu Glu Tyr Arg Ser Phe Cys Lys Ile Leu Phe Gln

Asp Tyr His Ser Phe Val Thr His Gly Cys Thr Val Asp Asn Pro Val 185

Leu Glu Arg Phe Ile Ser Leu Phe Asn Ser Val Ser Gln Trp Val Gln

Leu Met Ile Leu Ser Lys Pro Thr Ala Pro Gln Arg Ala Leu Val Ile 210 215

Thr His Phe Val His Val Ala Glu Lys Leu Leu Gln Leu Gln Asn Phe

Asn Thr Leu Met Ala Val Val Gly Gly Leu Ser His Ser Ser Ile Ser

		_		_,		C	vic	Val	Ser	Pro	Glu	Thr	Ile	Lys	Leu
			Glu 260					203							
		275	Leu				280					203			
	290		Arg			295					300				
Gly 305	Val	His	Leu	Lys	Asp 310	Leu	Val	Ala	Leu	Gln 315	Leu	Ala	Leu	Pro	Asp 320
Trp	Leu	Asp	Pro	Ala 325	Arg	Thr	Arg	Leu	Asn 330	Gly	Ala	Lys	Met	Lys 335	Gln
Leu	Phe	Ser	Ile 340	Leu	Glu	Glu	Leu	Ala 345	Met	Val	Thr	Ser	Leu 350	Arg	Pro
Pro	Val	Gln 355	Ala	Asn	Pro	Asp	Leu 360	Leu	Ser	Leu	Leu	Thr 365	Val	Ser	Leu
Asp	Gln 370	Tyr	Gln	Thr	Glu	Asp 375	Glu	Leu	Tyr	Gln	Leu 380	Ser	Leu	Gln	Arg
Glu 385	Pro	Arg	Ser	Lys	Ser 390	Ser	Pro	Thr	Ser	Pro 395	Thr	Ser	Cys	Thr	Pro 400
			Pro	405					410					310	
Lys	Leu	Asp	Gln 420	Ala	Leu	Val	Val	Glu 425	His	Ile	Glu	Lys	Met 430	Val	Glu
Ser	Val	Phe 435	Arg	Asn	Phe	Asp	Val 440	Asp	Gly	Asp	Gly	His 445	Ile	Ser	Gln
Glu	Glu 450	Phe	Gln	Ile	Ile	Arg 455	Gly	Asn	Phe	Pro	Tyr 460	Leu	Ser	Ala	Phe
Gly 465	Asp	Leu	Asp	Gln	Asn 470	Gln	Asp	Gly	Cys	11e 475	Ser	Arg	Glu	Glu	Met 480
Val	Ser	Tyr	Phe	Leu 485	Arg	Ser	Ser	Ser	Val 490	Leu	Gly	Gly	Arg	Met 495	Gly
Phe	Val	His	Asn 500	Phe	Gln	Glu	Ser	Asn 505	Ser	Leu	Arg	Pro	Val 510	Ala	CÃ2
Arg	His	Cys 515	Lys	Ala	Leu	Ile	Leu 520	Gly	Ile	Tyr	Lys	Gln 525	Gly	Leu	Lys
Cys	Arg 530	Ala	Cys	Gly	Val	Asn 535	Суѕ	His	Lys	Gln	Cys 540	Lys	Asp	Arg	Leu
Ser 545	Val	Glu	Cys	Arg	Arg 550		Ala	Gln	Ser	Val 555	Ser	Leu	Glu	Gly	Ser 560
Ala	Pro	Ser	Pro	Ser 565		Met	His	Ser	His 570	His	His	Arg	Ala	Phe 575	Ser
Phe	Ser	Leu	Pro 580	Arg	Pro	Gly	Arg	Arg 585	Gly	Ser	Arg	Pro	Pro 590	Glu	Ile
Arg	Glu	Glu 595	Glu	Val	Gln	Thr	Val 600	Glu	Asp	Gly	Val	Phe 605	Asp	Ile	His
Leu															

#### (2) INFORMATION FOR SEQ ID NO:8:

(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 832 base pairs(B) TYPE: nucleic acid

(C) STRANDEDNESS: single (D) TOPOLOGY: linear

#### (ii) MOLECULE TYPE: DNA

#### (ix) FEATURE:

(A) NAME/KEY: CDS
(B) LOCATION: 11..733

#### (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

GCC	CGCC									CGC Arg						49
							Leu					Ala			CGG	97
	Arg										Val				GCC Ala 45	145
					Lys					Ser					CTG Leu	193
										CTG Leu				Phe	GTG Val	241
										CGT Arg			Ser		CGC Arg	289
AGC Ser	TAT Tyr 95	GAT Asp	GAC Asp	CAG Gln	CTC Leu	CGC Arg 100	TCA Ser	GGT Gly	AGT Ser	CCC	CCA Pro 105	Lys	TCT Ser	CCA Pro	CGA Arg	337
ACC Thr 110	ACA Thr	GTC Val	CAT His	ĞAC Asp	AAG Lys 115	TCT	GCC Ala	CAC His	CAA Gln	ACA Thr 120	CAC His	AGC Ser	TCC Ser	TGG Trp	ACA Thr 125	385
CCC Pro	CCC Pro	AAC Asn	GCA Ala	CAG Gln 130	TAC Tyr	TGG Trp	TCC Ser	CAG Gln	TTT Phe 135	CAC His	AGC Ser	GTG Val	AGG Arg	CCA Pro 140	CAG Gln	433
										CAA Gln						481
GGG Gly	TAC Tyr	TGC Cys 160	CTC Leu	CTC Leu	CTC Leu	ATG Met	CTG Leu 165	GCG Ala	GGC Gly	ATG Met	Gly GGC	CTG Leu 170	CAC His	TAC Tyr	ATT Ile	529
GCC Ala	TTC Phe 175	AGG Arg	AAG Lys	GTG Val	AAG Lys	CAG Gln 180	ATG Met	CAC His	CTT Leu	AAC Asn	TTC Phe 185	ATG Met	GAT Asp	GAA Glu	AAG Lys	577

GAT Asp 190	Arg	G ATO	T ATC	C AC	A GCC Ala 195	Phe	TAC Ty:	c AAG	C GA n Gl	A GC u Al 20	a Ar	g GC. g Al	A CG	G GC0 g Ala	AGG A Arg 205
GCC Ala	AAC AST	AGA	GGG GL	210	e Lev	CAC Glr	G CAG	G GAG	G CG. u Arg 21	g Gl	A CGG	G CT	A GG	G CAC y Gli 220	G CGG n Arg
				Ser					n Gly					l Pro	CGG Arg
	_		Pro		GGG	GCTC	ACC	TGG	ATGG	GGC	TGC?	AGT (	CGT	rccc	GC .
TTT	TTTGCTTCCT TCCCTGGACG GCCCGCTCCC CGAAACGCGC GCAATAAAGT GATTCGCAG														
(2)	(2) INFORMATION FOR SEQ ID NO:9:  (i) SEQUENCE CHARACTERISTICS:														
		(i)	(A	) LE		: 24	1 am	ino		ls					
	-				POLO										
			MOLE SEOU			_			o ID	NO:	9.				•
Met 1	·	•	-							Arg		Trp	Pro	Arg 15	Asn
Pro	Pro	Ser	Arg 20	Leu	Leu	Gly	Ala	Ala 25		Gly	Gln	Arg	Ser 30	Arg	Pro
Ser	Thr	Tyr 35	Tyr	Glu	Leu	Leu	Gly 40	Val	His	Pro	Gly	Ala 45	Ser	Thr	Glu
Glu	Val 50	Lys	Arg	Ala	Phe	Phe 55	Ser	Lys	Ser	Lys	Glu 60	Leu	His	Pro	Asp
Arg 65	Asp	Pro	Gly	Asn	Pro 70	Ser	Leu	His	Ser	Arg 75	Phe	Val	Glu	Leu	Ser 80
Glu	Ala	Tyr	Arg	Val 85	Leu	Ser	Arg	Glu	Gln 90	Ser	Arg	Arg	Ser	Tyr 95	Asp
Asp	Gln	Leu	Arg 100	Ser	Gly	Ser	Pro	Pro 105	Lys	Ser	Pro	Arg	Thr 110	Thr	Val
His	Asp	Lys 115	Ser	Ala	His	Gln	Thr 120	His	Ser	Ser	Trp	Thr 125	Pro	Pro	Asn '
Ala	Gln 130	Tyr	Trp	Ser	Gln	Phe 135	His	Ser	Val	Arg	Pro 140	Gln	Gly	Pro	Gln
Leu 145	Arg	Gln	Gln	Gln	His 150	Lys	Gln	Asn	Lys	Gln 155	Val	Leu	Gly	Tyr	Cys 160
Leu	Leu	Leu	Met	Leu 165	Ala	Gly	Met	Gly	Leu 170	His	Tyr	Ile	Ala	Phe 175	Arg
Lys	Val	Lys	Gln 180	Met	His	Leu	Asn	Phe 1,85	Met	Asp	Glu	Lys	Asp 190	Arg	Ile
Ile	Thr	Ala	Phe	Туг	Asn	Glu	Ala	Arg	Ala	Arg	Ala	Arg	Ala	Asn	; Arg

210

195

200

Gly Ile Leu Gln Gln Glu Arg Gln Arg Leu Gly Gln Arg Gln Pro Pro

205

Pro Ser Glu Pro Thr Gln Gly Pro Glu Ile Val Pro Arg Gly Ala Gly 230 235 Pro (2) INFORMATION FOR SEQ ID NO:7: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 300 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPQLOGY: linear (ii) MOLECULE TYPE: DNA (ix) FEATURE: (A) NAME/KEY: CRS (B) LOCATION: 170 .300 (xi) SEQUENCE DESCRIPTION: SEO ID NO:7: CGATTTCATT CCTCGCTCCC CACAGGTCCC ACTCCCCAAA ATATTCCCAT CTTGTCCTAG 60 CCCATCCCCC AGACTATCTC AAGGACCAGC TGTBCGCACCG CCCCGACCT CCACTAGGCC 120 TGTGCCACCC GCTGCCTGCA GGAAGACGCC CGGTCCGGGG CCGGGTTAG CCC CAT 175 Pro His GGG AAC GGG GTT CGG TCC GAG CCC GGT GGG AGG CCC GAG CGC AGC 223 Gly Asn Gly Val Arg Ser Glu Pro Gly Gly Arg Led Pro Glu Arg Ser 10 CTG GGC CCA GCC CAC CCC GCG GCG GCC ATG GCA GGG ACC CTG GAC 271 Leu Gly Pro Ala His Pro Ala Pro Ala Ala Met Ala Gly Thr Leu Asp 20 25 CTG GAC AAG GGC TGC ACG GTG GAG GAG CT 300 Leu Asp Lys Gly Cys Thr Val Glu Glu Leu 35 (2) INFORMATION FOR SEQ ID NO:8: SEQUENCE CHARACTERISTICS: (A) LENGTH: 44 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear

SEQUENCE DESCRIPTION: SEQ ID NO.8:

his Gly Asn Gly Val Arg Ser Glu Pro Gly Gly Arg Leu Pro Glu

Bly Pro Ala His Pro Ala Pro Ala Ala Met Ala Giy Thr

Leu Asp Leu Asp Lys Gly Cys Thr Val Glu Glu Leu 35

- (2) INFORMATION FOR SEQ ID NO:9
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 15 base pairs
    - (B) TYPE: nucleig acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY! linear
  - (ii) MOLECULE TYPE: DNA

SEQUENCE DESCRIPTION: SEQ ID NO:9:

(2) INFORMATION FOR SEQ ID NO:10:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 13 amino acids

  - (B) TYPE: amino acid
    (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: Peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:10:

Asp Val Asp Glu Glu Asp Glu Val Glu Asp Ile Glu Phe

- (2) INFORMATION FOR SEQ ID NO:11:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 13 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:11:

Asp Val Asp Gly Asp Gly His Ile Ser Gln Glu Glu Phe

- (2) INFORMATION FOR SEQ ID NO:12:
  - (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 13 amino acids

- (B) TYPE: amino acid
- (C) STRANDEDNESS: single (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: Peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:12:

Asp His Asp Arg Asp Gly Phe Ile Ser Gln Glu Glu Phe

- (2) INFORMATION FOR SEQ ID NO:13:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 13 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:13:

Asp Gln Asn Gln Asp Gly Cys Ile Ser Arg Glu Glu Met

- (2) INFORMATION FOR SEQ ID NO:14:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 13 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:14:

Asp Val Asp Met Asp Gly Gln Ile Ser Lys Asp Glu Leu

- (2) INFORMATION FOR SEQ ID NO:15:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 37 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:15:

His Phe Val His Val Ala Glu Lys Leu Leu Gln Leu Gln Asn Phe Asn

Thr Leu Met Ala Val Val Gly Gly Leu Ser His Ser Ser Ile Ser Arg 20 25 30

Leu Lys Glu Thr His

- (2) INFORMATION FOR SEQ ID NO:16:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 37 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:16:

Lys Phe Val His Val Ala Lys His Leu Arg Lys Ile Asn Asn Phe Asn

Thr Leu Met Ser Val Val Gly Gly Ile Thr His Ser Ser Val Ala Arg

Leu Ala Lys Thr Tyr

- (2) INFORMATION FOR SEQ ID NO:17:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 amino acids

    - (B) TYPE: amino acid(C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:17:

His Asn Phe Gln Glu Ser Asn Ser Leu Arg Pro Val Ala Cys Arg His

Cys Lys Ala Leu Ile Leu Gly Ile Tyr Lys Gln Gly Leu Lys Cys Arg 20 25 30

Ala Cys Gly Val Asn Cys His Lys Gln Cys Lys Asp Arg Leu Ser Val

Glu Cys

- (2) INFORMATION FOR SEQ ID NO:18:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 50 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: Peptide
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:18:

His Asn Phe His Glu Thr Thr Phe Leu Thr Pro Thr Thr Cys Asn His

	1				5					10					15		
	Cys	Asn	Lys	Leu 20	Leu	Trp	Gly	Ile	Leu 25	Arg	Gln	Gly	Phe	Lys 30	Суѕ	Lys	
•	Asp	Cys	Gly 35	Leu	Ala	Val	His	Ser 40	Cys	Cys	Lys	Ser	Asn 45	Ala	Val	Ala	
	Glu	Суз 50		-		•											
(2)	INFOR	TAM	ON I	FOR S	SEQ I	א ס	):19:										
	(i)	(Ā) (B) (C)	JENCI LEN TYI STI TOI	IGTH: PE: r RANDE	: 15 nucle EDNES	base ic a S: s	pai cid ingl	rs				·					
	(ii)	MOLE	CULE	TYF	E: D	NA					•						
			٠													٠	
	(xi)	SEQU	JENCE	DES	CRIP	TION	: SE	Q ID	NO:	19:			,				
	ATCCCC												*		-		15
(2)	INFOR		•							•						•	
	(i) <sup>-</sup>	(A) (B) (C)	LENCE TYP STR TOP	GTH: E: n ANDE	21 ucle DNES	base ic a S: s	pai cid ingl	rs									
	(ii)	MOLE	CULE	TYP	E: D	NA:		•									
					,									,			
	(xi)	SEQU	ENCE	DES	CRIP	TION	: SE	Q ID	NO:	20:							
GAA'	PTCGGC.	A CG	AGCC	GACG	G												21
(2)	INFOR	MATI	ON F	OR S	EQ I	D NO	:21:						٠.				
	(i)	(A) (B) (C)	ENCE LEN TYP STR TOP	GTH: E: n' ANDE	78 ) ucle: DNES:	base ic a S: s:	pai: cid ingle	rs			-						
	(ii) 1	MOLE	CULE	TYP	E: Di	NA											
	(xi) :	SEQU	ENCE	DES	CRIP'	TION	: SE(	Q ID	NO:	21:							
ATG	GAGCAG	A AG	CTGA'	TCTC	CGA	GGAG	GAC (	TGC	cccc	G C	AGCT	GATO	CGC	CAGC	CAC		60
ccc	ccccc	G ČG	GCCA'	TG													78
(2)	INFOR	MATI	ON F	or s	EQ II	ON O	:22:			,							

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 26 amino acids
  - (B) TYPE: amino acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:22:

Met Glu Gln Lys Leu Ile Ser Glu Glu Asp Leu Pro Gly Ala Ala Gly

Ser Ala Ala His Pro Ala Pro Ala Ala Met 20

- (2) INFORMATION FOR SEQ ID NO:23:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 33 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:23:

GGATCCGCAG CCCACCCGC GCCGGCGGCC ATG

33

- (2) INFORMATION FOR SEQ ID NO:24:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 11 amino acids

    - (B) TYPE: amino acid(C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:24:

Gly Ser Ala Ala His Pro Ala Pro Ala Ala Met

- (2) INFORMATION FOR SEQ ID NO:25:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 21 base pairs

    - (B) TYPE: nucleic acid (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA

GTCATAGTCT GTCTCCTACT

(xi) SEQUENCE DESCRIPTION: SEQ ID NO	: 25 :
GGACAAAGTG TGTGATGAAC C	21
(2) INFORMATION FOR SEQ ID NO:26:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: DNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:	26:
CTCATCCTCC GTCTGATACT G	21
(2) INFORMATION FOR SEQ ID NO:27:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: DNA	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:	27:
GTAGATGTGG ATCAGCTTGG	20
(2) INFORMATION FOR SEQ ID NO:28:	
(i) SEQUENCE CHARACTERISTICS:     (A) LENGTH: 19 base pairs     (B) TYPE: nucleic acid     (C) STRANDEDNESS: single     (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: DNA	·
(xi) SEQUENCE DESCRIPTION: SEQ ID NO:	28:
AGGTGGAGAA TGGTCAAGG	19
(2) INFORMATION FOR SEQ ID NO:29:	
(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 20 base pairs  (B) TYPE: nucleic acid  (C) STRANDEDNESS: single  (D) TOPOLOGY: linear	
(ii) MOLECULE TYPE: DNA	
(xi) SECUENCE DESCRIPTION SEC ID NO.	20.

(2)	INFORMATION FOR SEQ ID NO:30:	
	<ul> <li>(i) SEQUENCE CHARACTERISTICS:</li> <li>(A) LENGTH: 20 base pairs</li> <li>(B) TYPE: nucleic acid</li> <li>(C) STRANDEDNESS: single</li> <li>(D) TOPOLOGY: linear</li> </ul>	
	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:30:	
ACA	TAGACAG CGTGCCTACC	20
(2)	INFORMATION FOR SEQ ID NO:31:	
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 20 base pairs  (B) TYPE: nucleic acid  (C) STRANDEDNESS: single  (D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:31:	
TAC	AACCTTA GGGACACCAG	20
(2)	INFORMATION FOR SEQ ID NO:32:	
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 20 base pairs  (B) TYPE: nucleic acid  (C) STRANDEDNESS: single  (D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:32:	
TGC	TGAGCCT GCTCACGGTG	20
(2)	INFORMATION FOR SEQ ID NO:33:	
	(i) SEQUENCE CHARACTERISTICS:  (A) LENGTH: 18 base pairs  (B) TYPE: nucleic acid  (C) STRANDEDNESS: single  (D) TOPOLOGY: linear	
	(ii) MOLECULE TYPE: DNA	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO:33:	
CAAC	GTGAACA GCACGTCC	18
	INFORMATION FOR SEQ ID NO:34:	
•	(i) SEQUENCE CHARACTERISTICS:	
	ini onformer emmeriment .	

18

	E)	LENGTH: 2: TYPE: nucl STRANDEDNI TOPOLOGY:	leic acid ESS: single	•				
	(ii) MOL	ECULE TYPE:	DNA					
			•					•
	(xi) SEQ	UENCE DESCRI	PTION: SEQ	ID NO:34:				
GAC	TATCTCA A	GGACCAGCT G					•	21
(2)	INFORMAT	ION FOR SEQ	ID NO:35:					
	(A (B (C	UENCE CHARAC ) LENGTH: 18 ) TYPE: nucl ) STRANDEDNE ) TOPOLOGY:	base pairs eic acid SS: single	5				
	(ii) MOL	ECULE TYPE:	DNA					
		· ·		,				
•	(xi) SEQ	UENCE DESCRI	PTION: SEQ	ID NO:35:				
GGT'	regeree G	AGCCCGG	٠					18
(2)	INFORMAT	ION FOR SEQ	ID NO:36:					
	(A) (B) (C)	JENCE CHARAC' LENGTH: 21 TYPE: nucl STRANDEDNE TOPOLOGY:	base pairs eic acid SS: single	·				
*	(ii) MOLE	CULE TYPE: 1	DNA			•		•
							,	
	(xi) SEQU	JENCE DESCRI	PTION: SEQ	ID NO:36:	-	•		
GGAG	CGATAC TO	CAAGTAGG T						21
(2)	INFORMATI	ON FOR SEQ	ID NO:37:					
-	(A) (B) (C)	TENCE CHARACT LENGTH: 18 TYPE: nucle STRANDEDNES TOPOLOGY: 1	base pairs ic acid SS: single					
	(ii) MOLE	CULE TYPE: I	ONA					

(2) INFORMATION FOR SEQ ID NO:38:

AGCGGGCCAG GCCCCTTC

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:37:

WO 98/53061 PCT/AU98/00380

- 83 -

	(i)	SEQUENCE CHARACTERISTICS:  (A) LENGTH: 20 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear			·	
	(ii)	MOLECULE TYPE: DNA	•	•		
	(xi)	SEQUENCE DESCRIPTION: SEQ ID	NO:38:			
CA	rcctgg	TC CAATGCGCTC	•			20
(2)	INFO	RMATION FOR SEQ ID NO:39:	to some			
	(i)	SEQUENCE CHARACTERISTICS: (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear				. `
	(ii)	MOLECULE TYPE: DNA				
	(xi)	SEQUENCE DESCRIPTION: SEQ ID	NO:39:	4		
GCA	CTGAG	GA AGTTAAACGA GC				22
•						
(2)	INFOR	RMATION FOR SEQ ID NO:40:				
	(i)	SEQUENCE CHARACTERISTICS:  (A) LENGTH: 22 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear				
	(ii)	MOLECULE TYPE: DNA				
	(xi)	SEQUENCE DESCRIPTION: SEQ ID	NO:40:			
GCT	GTTTA	A CTTCCTCAGT GC				22
(2)	INFOR	MATION FOR SEQ ID NO:41:				
	(i)	SEQUENCE CHARACTERISTICS:  (A) LENGTH: 21 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear				-

(ii) MOLECULE TYPE: DNA

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:41:

GCTCAGCTCC ACAAAGCGGC T

(2) INFORMATION FOR SEQ ID NO:42:

(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 19 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (xi) SEQUENCE DESCRIPTION: SEQ ID NO:42: ACCAGCTCCG CTCAGGTAG

19

- (2) INFORMATION FOR SEQ ID NO:43:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 20 base pairs
    - (B) TYPE: nucleic acid (C) STRANDEDNESS: single

    - (D) TOPOLOGY: linear.
  - (ii) MOLECULE TYPE: DNA
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:43:

#### TCCAGGAGCT GTGTGTTTGG

20

- (2) INFORMATION FOR SEQ ID NO:44:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 19 base pairs
    - (B) TYPE: nucleic acid
    - (C) STRANDEDNESS: single
      (D) TOPOLOGY: linear

    - (ii) MOLECULE TYPE: DNA
    - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:44:

#### CCAGTTTCAC AGCGTGAGG

19

- (2) INFORMATION FOR SEQ ID NO:45:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 19 base pairs

    - (B) TYPE: nucleic acid (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: DNA
  - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:45:

#### CAGCATGAGG AGGAGGCAG

19

Please and SEQIB: 46-110 as Set forth in the substitute ? I Sequence Listing.



#### **BRIEF DESCRIPTION OF THE FIGURES**

Figure 1 is a representation of the nucleotide sequence (SEQ ID NO: 2) and corresponding amino acid sequence (SEQ ID NO: 3) of *mcg*4.

Figure 2 is a representation of the alignment of the human MCG4 amino acid sequence (amino acids: 1-60, SEQ ID NO: 46) (Query) with a translation of a partial murine expressed sequence tag (EST) (SEQ ID NO: 47) (Subject).

Figure 3 is a representation of the alignment of the human MCG4 amino acid sequence (amino acids 7-109; SEQ ID NO: 48) (Query) with a translation of a partial nematode EST (SEQ ID NO: 49) (subject).

Figure 4 is a diagrammatic representation showing a predicted structure of MCG4 where H and C represent histidine and cysteine residues, respectively and X refers to any amino acid residue. Zn represents zinc atoms.

Figure 5 is a representation of sensitive sequence homology search of related cysteine-containing motifs in another *Caenorhabditis elegans* protein. Query: amino acids 56-100 (SEQ ID NO: 50). Subject: SEQ ID NO: 51.

Figure 6 is a representation showing that a related cysteine containing motif is present in the GATA-binding transcription factor from *Sacchromyces pombe*. Queries: amino acids 35-58 (SEQ ID NO: 52); amino acids 162-227 (SEQ ID NO: 54). Subjects: amino acids 175-198 (SEQ ID NO: 53); amino acids 441-506 (SEQ ID NO: 55).

Figure 7 is a Northern blot showing expression of *mcg*4 in various cultured human cancer cell lines. Lanes 1-5, respectively, represent the hybridization signal from 15 μg total RNA derived from various human cancer cells lines. Lanes 1-5, respectively, contain RNA from H69 lung carcinoma cells, JAM ovary carcinoma cells, BT20 breast carcinoma cells, HaCat transformed keratinocytes, T24 bladder carcinoma cells.

Figure 8 is a representation of a partial alignment of *mcg*4 with human ESTs AA074703 and AA134788. Queries: nucleotides 446-704 (SEQ ID NO: 56); nucleotides 398-452 (SEQ ID NO: 58); nucleotides 767-810 (SEQ ID NO: 60); nucleotides 731-765 (SEQ ID NO: 62); nucleotides 701-732 (SEQ ID NO: 64); nucleotides 498-687 (SEQ ID NO: 66); nucleotides 398-495 (SEQ ID NO: 68); nucleotides 702-761 (SEQ ID NO: 70). Subjects: nucleotides 49-307 (SEQ ID NO: 57); nucleotides 2-56 (SEQ ID NO: 59); nucleotides 373-416 (SEQ ID NO: 61);



nucleotides 336-370 (SEQ ID NO: 63); nucleotides 305-336 (SEQ ID NO: 65); nucleotides 103-292 (SEQ ID NO: 67); nucleotides 2-99 (SEQ ID NO: 69); nucleotides 309-368 (SEQ ID NO: 1).

Figure 9 is a representative of the partial nucleotide sequence alignment between a human (W32939) (SEQ ID NO: 72) and mouse (AA242159) (SEQ ID NO: 73) *mcg*4 –like EST in the putative 5' UTR of the *mcg*4 cDNA. The putative initiation codon is underlined and the region upstream represents 5' UTR.

Figure 10 is a representation showing MacVector alignment of MCG4 with forward translations of ESTs AA134788 and AA074703. The nucleotide sequences are shown in Figure 8.

Figure 11 is a diagrammatic representation of the domains of MCG4 zinc finger consensus: CX<sub>2</sub>HX<sub>4</sub>CX<sub>2</sub>CX<sub>4</sub>HX<sub>2</sub>CX<sub>17</sub>CX<sub>2</sub>CX<sub>18</sub>HX<sub>2</sub>CX<sub>18</sub>CX<sub>2</sub>C (SEQ ID NO: 74); acidic domain consensus: 9/34 amino acids negatively charged, 0/34 positively charged; basic domain consensus: 13/55 amino acids positively charged, 0/55 negatively charged; leucine zipper domain consensus: LX<sub>6</sub>LX<sub>6</sub>RX<sub>6</sub>LX<sub>6</sub>L (SEQ ID NO: 75); alternate "novel" leucine zipper-like motif where leucine would not be aligned along the one surface to an alpha helix domain: (aa261) LX<sub>6</sub>LXLX<sub>6</sub>LXLX<sub>6</sub>L (aa286) (SEQ ID NO: 76).

Figure 12 is a representation showing similarity of MCG7 WITH GEFs of various organisms.

Figure 13(a) is a representation of the nucleotide sequence (SEQ ID NO: 4) and corresponding amino acid sequence (SEQ ID NO: 5) of *mcg7*. Nucleotides 183-288 are an alternative spliced exon (shown in lower case).

Figure 13(b) is a representation of the partial nucleotide sequence (SEQ ID NO: 6) and corresponding amino acid sequence (SEQ ID NO: 7) OF *mcg7* but without the exon shown in Fig. 13(a). Amino acids have been numbered from the first methionine codon (underlined). The cDNA molecules of Fig. 13(a) and Fig. 13(b) differ by the inclusion and exclusion of the exon of nucleotides 183-288.

Figure 14 is a representation showing a comparison between MCG7 (SEQ ID NO: 7) and a homologue from *Caenorhabditis elegans* (SEQ ID NO: 77) using the BESTFIT algorithm. In the figure, the following sequences are underlined:

#### EF-Hand=PROSITE DATABASE NO. PD0C00018

la nematode DVDEEDEVEDIEF (SEQ ID NO: 10)

lb human DVDGDGHISQEEF (SEQ ID NO: 11)

nematode DHDRDGFISQEEF (SEQ ID NO: 12)

lc human DQNQDGCISREEM (SEQ ID NO: 13)

nematode DVDMDGQISKDEL (SEQ ID NO: 14)

#### GUANINE NT BINDING REGION = BLOCKS DATABASE NO. BL00720B

2 human HFVHVAEKLLQLQNFNTLMAVVGGLSHSSISRLKETH (SEQ ID NO: 15) nematode KFVHVAKHLRKINNFNTLMSVVGGITHSSVARLAKTY (SEQ ID NO: 16)

#### **DaG-PE BINDING DOMAIN = PROSITE DATABASE NO. PD0C00379**

3 human HNFQESNSLRPVACRHCKALILGIYKQGLKCRACGVNCHKQCKDRLSVEC (SEQ ID NO: 17)

nematode HNFHETTFLTPTTCNHCNKLLWGILRQGFKCKDCGLAVHSCCKSNAVAEC (SEQ ID NO: 18)

Figure 15 is a representation of an alignment of human and a partial (5' UTR and partial coding sequence) murine mcg7 cDNA (GenBank Acc.No. W71787 and AA237373). The putative initiation codon is underlined. The murine sequence (SEQ ID NO: 78) represents a composite of 2 partial cDNA sequences from the EST database (accession numbers W71787 and AA237373). Nucleotide differences between human and murine sequences are shown in lower case lettering and identical residues are indicated with asterisks.

Figure 16 is a representation of further 5' nucleotide (SEQ ID NO: 79) and corresponding amino acid sequence (SEQ ID NO: 80) human *mcg*7. Nucleotide positions 1-321 were derived from GenBank Acc. No. AC000134 and nucleotides 322 onwards from Figure 13(a). Two inframe initiation codons are underlined. Asterisks denote in-frame stop codons.

Figure 17 is a graphical representation of a GDP release assay. □Experiment #1 (mean of duplicate). ◇Experiment #2 (mean of duplicates). The exchange reaction contained 36pmols of GST-MCG (N-terminally truncated; encoded by Construct B in Fig. 18) and 1.6-12.8 pmols of recombinant GST-N-Ras.GDP. Reaction time 6 min.

Estimated reaction constants:

 $K_m = 2.1 \mu M$ ,  $V_{max} = 37 \text{ pMol/6min/36pMol [Expt.#1]}$ 

 $K_m = 1.5 \mu M$ ,  $V_{max} = 30.3 p Mol/6 min/36 p Mol[Expt.#2]$ 

Figure 18 depicts various recombinant plasmids containing partial or full-length mcg7.

Figure 19 is a representation of the nucleotide sequence (SEQ ID NO: 8) and corresponding amino acid sequence (SEQ ID NO: 9) of *mcg18*.

Figure 20 is representation showing that MCG18 (amino acids 35-96, SEQ ID NO: 81] (Query) has partial homology to *E. coli* DnaJ (amino acids 6-65, SEQ ID NO: 82) (Subject).

Figure 21 is a representation showing that MCG18 has homotology to two *Caenorhabitis elegans* proteins. Queries: amino acids 28-64 (SEQ ID NO: 83); amino acids 71-102 (SEQ ID NO: 85); amino acids 162-203 (SEQ ID NO: 87); amino acids 35-75 (SEQ ID NO: 89); amino acids 75-96 (SEQ ID NO:91); amino acids 141-184 (SEQ ID NO: 93). Subjects: amino acids 22-58 (SEQ ID NO:84); amino acids 64-95 (SEQ ID NO: 86); amino acids 158-199 (SEQ ID NO: 88); amino acids 19-50 (SEQ ID NO: 90); amino acids 71-92 (SEQ ID NO: 92); amino acids 145-188 (SEQ ID NO: 94).

Figure 22 is a representation showing that MCG18 has homology to a *Saccharomyces pombe* protein. Queries: amino acids 35-70 (SEQ ID NO: 95); amino acids 75-114 (SEQ ID NO: 97); amino acids 190-218 (SEQ ID NO: 99); amino acids 140-158 (SEQ ID NO: 101). Subjects: amino acids 9-44 (SEQ ID NO: 96); amino acids 50-89 (SEQ ID NO: 98); amino acids 149-177 (SEQ ID NO:100); amino acids 44-62 (SEQ ID NO: 102).

Figure 23 is a representation showing homology of MCG18 to a *Drosophila virilis* protein. Queries: amino acids 26-95 (SEQ ID NO: 103). Subject: amino acids 72-142 (SEQ ID NO: 104).

Figure 24 is a representation showing homology of MCG18 to human DnaJ protein HDJ-2/HSDJ (SEQ ID NO: 105), HDJ-1/HSP40 (SEQ ID NO: 106) and HSJ1 (SEQ ID NO: 107).

Figure 25 is a representation of the nucleotide (SEQ ID NO: 108) and corresponding amino acid sequence (SEQ ID NO: 109) of murine *mcg18*.

Figure 26 is a representation of homology between human and murine MCG18 (SEQ ID NO: 9 and SEQ ID NO: 109).

Figure 27 depicts nucleotide sequence (SEQ ID NO: 110) corresponding to the 5' untranslated region of human *mcg18*.

Figure 28 depicts a Northern blot showing expression of mcg 18 transcripts in total RNA isolated from various human cancer cell lines grown in culture. Lanes 1-5 respectively contain 15µg RNA from H69 lung carcinoma cells, JAM ovary carcinoma cells, BT20 breast carcinoma cells, HaCat transformed keratinocytes, T24 bladder carcinoma cells.